



Please type a plus sign (+) in this box

PTO/SB (12-97)
 OMB 0651-0031

Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE
 Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it displays a valid OMB control number.

Modified Form 1449/PTO SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT BY APPLICANT (use as many sheets as necessary)		Application Number	10/692,523
		Filing Date	10/24/03
		First Named Inventor	Bacopoulos
		Group Art Unit / Confirmation No.	1614 / 9840
		Examiner Name	Cybille Delacroix-Muirheid
		Attorney Docket Number	24852-513

U.S. PATENT DOCUMENTS							
Exam Initials	Cite No.	U.S. Patent Document No.	Issue Date	Name of Patentee(s) or Applicant(s)	Class	Sub Class	Filing Date
JDA	A23*	4,690,918	09/01/87	Beppu, et al.			
JDA	A24*	5,654,333	08/05/97	The United States of America as represented by the Department of Health and Human Services			
JDA	A25*	6,239,176	05/29/01	Beacon Laboratories, Inc. et al.			
JDA	A26*	6,262,116	07/17/01	Sloan-Kettering Institute for Cancer Research			
JDA	A27*	6,451,334	09/17/02	Perrine			
JDA	A28*	6,495,719	12/17/02	CircaGen Pharmaceutical			
JDA	A29*	2003/0114525	06/19/03	Kammer, et al.			
JDA	A30*	2004/0132643	07/08/04	Fojo, et al.			
JDA	A31*	2004/0167184	08/26/04	Wlech, et al.			
JDA	A32	6,495,719	12/17/02	Lan-Harget, et al.			

FOREIGN PATENT DOCUMENTS						
Exam Initials	Cite No.	Foreign Patent Document Office	Document Number	Name of Patentee(s) or Applicant(s)	Date of Publication	English Yes No
JDA	B12*	WO	98/39965	Beacon Laboratories, LLC	09/17/98	X
JDA	B13*	WO	02/15921	The Government of the United States of America	02/28/02	X
JDA	B14*	WO	02/055017	Wake Forest University	07/18/02	X

NON PATENT LITERATURE DOCUMENTS		
Exam Initials	Cite No.	Name of Author, Title (when appropriate), Publication, Volume, Page(s), Date, Etc.
JDA	C83*	"Aton Pharma, Inc. Announces Initiation of Two Phase II Trials to Evaluate Efficacy of HDAC Inhibitor SAHA", October 30, 2002.
JDA	C84*	"Aton Pharma, Inc. Announces Phase I Clinical Trial of SAHA in Advanced Leukemias", July 1, 2003.
JDA	C85*	"Aton Pharma, Inc. Appoints Judy H. Chiao, M.D., as Vice President, Oncology Clinical Research and Development", September 20, 2002.
JDA	C86*	"Aton Pharma, Inc. Presents Phase I Trial Data of Anti-Cancer Agent SAHA in Patients with hematological Malignancy at ASCO", June 2, 2003.
JDA	C87*	"Aton Pharma, Inc. Presents Phase I Trial Data on Anti-Cancer Agent SAHA at EORTC/NCI/AACR Symposium", November 21, 2002.
JDA	C88*	"Aton Pharma, Inc. Received Orphan Drug Designation for SAHA in Multiple Myeloma and Initiates Phase I Trial", October 13, 2003.

NON PATENT LITERATURE DOCUMENTS		
Exam. Initials	Cite No.	Name of Author, Title (when appropriate), Publication, Volume, Page(s), Date, Etc.
JDA	C89*	"Aton Pharma, Inc. Reports on Phase I Trial of SAHA", August 14, 2002.
	C90*	Adhikari, D et al., Proceedings of the American Association for Cancer Research Annual Meeting, (1998), Vol. 39, p 312, "Radiosensitization of Lymphoma Cell Lines by Sodium Butyrate".
	C91*	Alexandrov, I et al., FEBS Letters, (1998), Vol. 434, pp 209-214, "Sodium Butyrate Suppresses Apoptosis in Human Burkitt Lymphomas and Murine Plasmacytomas Bearing c-myc Translocations".
	C92*	Almenara, J et al., Leukemia (2002), Vol. 16, pp 1331-1343, "Synergistic Induction of Mitochondrial Damage and Apoptosis in Human Leukemia Cells by Flavopiridol and the Histone Deacetylase Inhibitor Suberoylanilide Hydroxamic Acid (SAHA)".
	C93*	Amin HM et al., British Journal of Haematology (2001), Vol. 115, pp 287-297, "Histone Deacetylase Inhibitors Induce Caspase-Dependent Apoptosis and Downregulation of Daxx in Acute Promyelocytic Leukaemia with t(15;17)".
	C94*	Aron, JL et al., Blood (2003), Vol. 102, No. 2, pp 652-658, "Depsipeptide (FR901228) Induces Histone Acetylation and Inhibition of Histone Deacetylase in Chronic Lymphocytic Leukemia Cells Concurrent With Activation of Caspase 8-mediated Apoptosis and Down-Regulation of c-FLIP Protein".
	C95*	Benoit, NE et al., Immunopharmacology, (1996), Vol. 35, pp 129-139, "Increased inhibition of Proliferation of Human B Cell Lymphomas Following Ligitation of CD40, and Either CD19, CD20, CD95 or Surface Immunoglobulin".
	C96*	Bode, J et al., Journal of Interferon Research, (1982), Vol. 2, No. 2, pp 159-166, "Links Between Effects of Butyrate on Histone Hyperacetylation and Regulation of Interferon Synthesis in Namalva and FS-4 Cell Lines".
	C97*	Buckley, AR et al., Cell Growth & Differentiation (1996), Vol. 7, pp 1713-1721, "Alterations in pim-1 and c-myc Expression Associated with Sodium Butyrate-induced Growth Factor Dependency in Autonomous Rat Nb2 Lymphoma Cells".
	C98*	Buckley, AR et al., Proceedings of the American Association for Cancer Research Annual Meeting, (1997), Vol. 38, p 193, "Reversal of Apoptosis Resistance by Butyrate in rat Nb2 Lymphoma Cells".
	C99*	Byrd, JC et al., Blood (1999), Vol. 94, No. 4, pp 1401-1408, "Depsipeptide (FR901228): A Novel Therapeutic Agent with Selective, In Vitro Activity Against Human B-Cell Chronic Lymphocytic Leukemia Cells".
	C100	Cao, et al. (2001), Am. J. Respir. Cell Mol. Biol., 25:562-8, "Histone Deacetylase Inhibitor Downregulation of bcl-xl Gene Expression Leads to Apoptotic Cell Death in Mesothelioma".
	C101*	Carducci, MA et al., Clinical Cancer Research (2001), Vol. 7, No. 10, pp 3047-3055, "A Phase I Clinical and Pharmacological Evaluation of Sodium Phenylbutyrate on an 120-h Infusion Schedule".
	C102*	Dear, AE et al., Biochimica et Biophysica Acta, (2000), Vol. 1492, pp 15-22, "The Novel Anti-Tumour Agent Oxfamifatin Differentially Regulates Urokinase and Plasminogen Activator Inhibitor Type 2 Expression and Inhibits Urokinase-Mediated Proteolytic Activity".
	C103*	Desai, D et al., Anticancer Research (2003), Vol. 23, pp 499-504, "Chemopreventive Efficacy of Suberoylanilide Hydroxamic Acid (SAHA) Against 4-(Methylnitrosamino)-1-(3-pyridyl)-1-butanone (NNK)-induced Lung Tumorigenesis in Female A/J Mice".
	C104*	Dhordain, P et al., Nucleic Acids Research, (1998), Vol. 26, No. 20, pp 4645-4651, "The LAZ3(BCL-6) Oncoprotein Recruits a SMRT/mSIN3A/Histone Deacetylase Containing Complex to Mediate Transcriptional Repression".
	C105*	Edelman, MJ et al., Cancer Chemotherapy and Pharmacology (2003), Vol. 51, pp 439-444, "Clinical and Pharmacologic Study of Tributyryl: An Oral Butyrate Prodrug".
	C106*	Feinman, R et al., Blood (2002), Vol. 100, No. 11, pp Abstract 3195, "The Histone Deacetylase Inhibitor, Suberoylanilide Hydroxamic Acid, Induces Apoptosis of Multiple Myeloma Cells".
	C107*	Filippovich, I et al., Biochemical and Biophysical Research Communications, (1994), Vol. 198, pp 257-265, "Butyrate Induced Apoptosis in Lymphoid Cells Preceded by Transient Over-Expression of HSP70 mRNA".
	C108*	Foss, FM et al., Blood, (1993), Vol. 82, No. 10, Suppl. 1, p 564A, "Biomodulatory Effects of Butyric Acid Derivatives on Leukemia and Lymphoma Cells".
↓	C109	Gediya, et al., J. Med. Chem. (2005), Vol. 48, pp 5047-5051, "A New Simple and High-Yield Synthesis of Suberoylanilide Hydroxamic Acid and Its Inhibitory Effect Alone or in Combination with Retinoids on Proliferation of Human Prostate Cancer Cells".
JDA	C110*	Gelmetti, V et al., Molecular and Cellular Biology (1998), Vol. 18, No. 12, pp 7185-7191, "Aberrant Recruitment of the Nuclear Receptor Corepressor-Histone Deacetylase Complex by the Acute Myeloid Leukemia Fusion Partner ETO".

NON PATENT LITERATURE DOCUMENTS		
Exam Initials	Cite No.	Name of Author, Title (when appropriate), Publication, Volume, Page(s), Date, Etc.
JDA	C111*	Gerbitz, A, Oncogene, (1999), Vol. 18, pp 1745-1753, "Deregulation of the Proto-Oncogene c-myc Through t(8;22) Translocation in Burkitt's Lymphoma".
	C112*	Gilbert, J et al., Clinical Cancer Research (2001), Vol. 7, No. 8, pp 2292-2300, "A Phase I Dose Escalation and Bioavailability Study of Oral Sodium Phenylbutyrate in Patients with Refractory Solid Tumor Malignancies".
	C113*	Grisolano, JL et al., Proceedings of the National Academy of Sciences (2003), Vol. 100, No. 16, pp 9506-9511, "An Activated Receptor Tyrosine Kinase, TEL/PDGFBetaR, Cooperates with AML1/ETO to Induce Acute Myeloid Leukemia in Mice".
	C114*	Harris, NL et al., Blood (1994), Vol. 84, No. 5, pp 1361-1392, "A Revised European-American Classification of Lymphoid Neoplasms: A Proposal From the International Lymphoma Study Group".
	C115*	Jaboin, J et al., Cancer Research (2002), Vol. 62, No. 21, pp 6108-6115, "MS-27-275, an Inhibitor of Histone Deacetylase, Has Marked in Vitro and in Vivo Antitumor Activity against Pediatric Solid Tumors".
	C116*	Kurita-Ochiai, T et al., Infection and Immunity, (1998), Vol. 66, No. 6, pp 2587-2594, "Volatile Fatty Acid, Metabolic By-Product of Periodontopathic Bacteria, Induces Apoptosis in WEHI 231 and RAJI B Lymphoma Cells and Splenic B Cells".
	C117*	Liu, Z et al., Journal of Cancer Research and Clinical Oncology, (1998), Vol. 124, pp 541-548, "Synergistic Effect of Epstein-Barr Virus and Tumor Promoters on Induction of Lymphoma and Carcinoma in Nude Mice".
	C118*	Madisen, L et al., Molecular and Cellular Biology, (1998), Vol. 18, No. 11, pp 6281-6292, "The Immunoglobulin Heavy Chain Locus Control Region Increases Histone Acetylation along Linked c-myc Genes".
	C119*	Niitsu, N et al., Molecular Pharmacology, (2000), Vol. 58, pp 27-36, "Anticancer Derivative of Butyric Acid (Pivalyloxymethyl Butyrate) Specifically Potentiates the Cytotoxicity of Doxorubicin and Daunorubicin Through the Suppression of Microsomal Glycosidic Activity".
	C120*	Orr, D et al., 2000 ASCO Annual Meeting, Abstract No. 763, "Phase I Pharmacokinetic (PK) Study of CI-994 in Combination with Gemcitabine (GEM) in Patients with Advanced Solid Tumors".
	C121*	Polack, A et al., The EMBO Journal, (1993), Vol. 12, No. 10, pp 3913-3920, "Regulatory Elements in the Immunoglobulin Kappa Locus Induce c-myc Activation and the Promoter Shift in Burkitt's Lymphoma Cells".
	C122*	Rezuke, WN et al., Clinical Chemistry (1997), Vol. 43, No. 10, pp 1814-1823, "Molecular Diagnosis of B- and T-cell Lymphomas: Fundamental Principles and Clinical Applications".
	C123*	Rottlieb, C et al., International Journal of Cancer, (1995), Vol. 62, pp 697-702, "Among 17 Inducers of Differentiation Only Sodium Butyrate Causes a Permanent Down-Regulation of c-myc in Burkitt's Lymphoma".
	C124*	Rottlieb, C et al., International Journal of Cancer, (1996), Vol. 67, pp 724-729, "Structure-Activity Relationship of 17 Structural Analogues of N-Butyric Acid Upon c-myc Expression".
	C125*	Rubio, MA et al., Blood, (1995), Vol. 86, No. 10, pp 3715-3724, "Granulocyte-Macrophage Colony-Stimulating Factor, Phorbol Ester, and Sodium Butyrate Induce the CD11c Integrin Gene Promoter Activity During Myeloid Cell Differentiation".
	C126*	Schrump, DS et al., Clinical Lung Cancer (2002), Vol. 4, No. 3, pp 186-192, "Phase I Study of Sequential Deoxyazacytidine/depsipeptide Infusion in Patients with Malignancies Involving Lungs or Pleura".
	C127*	Vrana JA et al., Oncogene 1999), Vol. 18, pp 7016-7025, "Induction of Apoptosis in U937 Human Leukemia Cells by Suberoylanilide Hydroxamic Acid (SAHA) Proceeds Through Pathways That are Regulated by Bcl-2/Bcl-XL, c-Jun, and p21CIP1, but independent of p53".
	C128	Waheed et al. (2000), Proceedings of the American Association for Cancer Research Meeting, (91st, San Francisco, 41:808, Abstract 5135, "The Histone Deacetylase Inhibitor FR 901228 Induces SV40T/T Antigen Expression and P53 Hyperacetylation in Human Pleural Mesothelioma Cells".
	C129*	Watanabe, M et al., Cancer Research (1990), Vol. 50, pp 3245-3248, "Effect of liposomes containing sodium butyrate conjugated with anti-CD19 monoclonal antibody on in vitro and in vivo growth of malignant lymphoma".
	C130	Weiser et al. (2001), J. Immunotherapy, 24:151-61, "Sequential 5-Aza-2'deoxyctidine-Depsipeptide FR901228 Treatment Induces Apoptosis Preferentially in Cancer Cells and Facilitates Their Recognition by Cytolytic T Lymphocytes Specific for NY-ESO-1".
	C131*	Yu, C et al., Cancer Research (2001), Vol. 63, pp 2118-2126, "Histone Deacetylase Inhibitors Promote ST1571-Mediated Apoptosis in ST1571-Sensitive and -Resistant Bcr/Abl+ Human Myeloid Leukemia Cells".
↓	C132*	Zhang, M et al., Cell Stress & Chaperones, (1998), Vol 3, No. 1, pp 57-66, "Heat-Induced Proteolysis of HSF Causes Premature Deactivation of the Heat Shock Response in Nb2 Lymphoma Cells".
JDA	C133	Bruner, RJ et al., Blood (2002), 44th Annual Meeting of the American Society of Hematology, Vol. 100, No. 11, pp Abstract No. 1492, "Phase I trial of the histone deacetylase inhibitor depsipeptide (FR901228) in fludarabine

NON PATENT LITERATURE DOCUMENTS		
Exam Initials	Cite No.	Name of Author, Title (when appropriate), Publication, Volume, Page(s), Date, Etc.
		refractory chronic lymphocytic leukemia".
JDA	C134	Guo, F et al., American Society of Hematology, (December 6-10, 2002), p 268b, Abstract 4602 "Co-treatment with the histone deacetylase inhibitor suberoylanilide hydroxamic acid (SAHA) enhances Apo-2L/TRAIL-induced death inducing signaling complex and apoptosis of human acute lymphoid leukemia cells".
JDA	C135	Heaney, M et al., 2003 ASCO Annual Meeting, Proceedings of the American Society of Clinical Oncology, (2003) Vol. 22, p 577, Abstract 2321, "Clinical experience with the histone deacetylase (HDAC) inhibitor suberoylanilide hydroxamic acid (SAHA) in heavily pre-treated patients with hematological malignancies".
JDA	C136	Marcucci, G et al., Blood, (2002), 44th Annual Meeting of the American Society of Hematology", Vol. 100, No. 11, pp Abstract No. 317, "Phase I trial of the histone deacetylase inhibitor depsipeptide (FR901228) in acute myeloid leukemia (AML)".
JDA	C137	Nimmanapalli, R et al., American Society of Hematology, (December 6-10, 2002). 14 pages, "Co-treatment with the histone deacetylase inhibitor suberoylanilide hydroxamic acid (SAHA) enhances Gleevee-induced apoptosis of Bcr-Abl positive human acute leukemia cells".
JDA	C138	Nimmanapalli, R et al., Blood (2003), Vol. 101, No. 8, pp 3236-3239, "Cotreatment with the histone deacetylase inhibitor suberoylanilide hydroxamic acid (SAHA) enhances imatinib-induced apoptosis of Bcr-Abl-positive human acute leukemia cells".
JDA	C139	Tabe, Y et al., Blood (2002), 44th Annual Meeting of the American Society of Hematology, Vol. 100, No. 11, pp Abstract No. 3028, "Effects of histone deacetylase inhibitor suberoylanilide hydroxamic acid (SAHA) and DNA methylation inhibitor 5-aza-2'-deoxycytidine (DAC) on the transcriptional activation of RARbeta and p21WAF in acute promyelocytic leukemia cells".
JDA	C140	Zhang, C et al, The Journal of Investigative Dermatology (2003), Vol. 121, No. 1, pp Abstract 1189, "The histone inhibitor suberoylanilide hydroxamic acid induces apoptosis in cutaneous T cell lymphoma cells".

*a copy of this reference is not provided as it was previously cited by or submitted to the office in a prior application, Serial No. 10/650,025, filed August 26, 2003, and relied upon for an earlier filing date under 35 U.S.C. §120 (continuation, continuation-in-part, and divisional applications).

Examiner Signature	/James Anderson/	Date Considered	12/21/2006
--------------------	------------------	-----------------	------------

EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.